

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In Re Patent Application Of:
Kipp et al.

For: MICROPRECIPITATION
METHOD FOR PREPARING
SUBMICRON SUSPENSIONS

Serial No.: 09/874,637

Filed: June 5, 2001

Examiner: Simon J. Oh

Art Unit: 1615

Conf. No. 5102

CERTIFICATE OF MAILING

I hereby certify that this paper is being deposited with the United States Postal Office with sufficient postage as first class mail in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450 on August 9, 2004.

Judith L. Goldberg
Judith L. Goldberg

DECLARATION

Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22202

Sir:

I, Mark Doty aver as follows:

1. I am over the age of twenty-one years and make these statements from my own personal knowledge.

2. I have received a Bachelors Degree in Chemical Engineering from the University of Minnesota in 1987 and a Doctor of Philosophy Degree in Chemistry from Iowa State University in 1995.

3. I have been employed by Baxter since 1995 and currently hold the position of Associate Director of Research.

4. I am a joint inventor on U.S. Patent Application Serial No. 09/874,637.

5. I have reviewed the claims of this application and I believe that the claimed invention was conceived prior to January 7, 2000 and I have diligently worked on this project since this time.

6. As evidence of this I have attached laboratory notebook pages with dates redacted. The laboratory notebook pages are not necessarily the earliest date of conception of

the claimed invention and do not represent the results of an exhaustive search for evidence of the earliest date of conception or reduction to practice.

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true, and further, I acknowledge that willful false statements and the like are punishable by fine or imprisonment, or both, under §1001 of Title 18 of the United States Code and may jeopardize the validity of the application or any patent issuing thereon.

Date: August 9, 2004

BY


Mark J. Doty

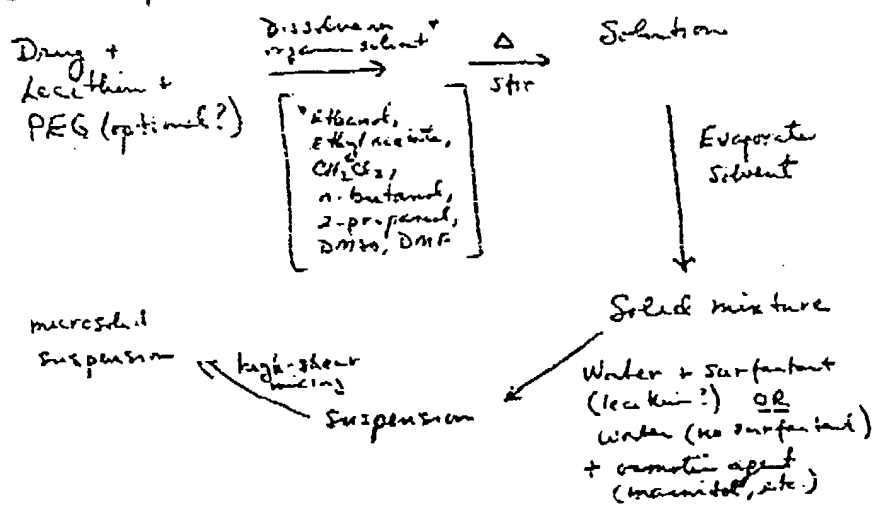
BOOK NO. 20744 - 39

Solid Dispersion of Prednisolone - Lecithin

FPM 170

Notebook

Purpose: Test the hypothesis that one can make nanocapsules (or minimize particle size in a microsuspension) by first preparing a solid dispersion of the drug in a matrix such as PEG (polyethylene glycol) and/or phosphatidylglycerol^{2, 3, 4, 5} and/or phosphatidyl choline (or lecithin). The solid is then resuspended in an aqueous solvent (water + surfactant). Hopefully, high shear mixing (stirring? homogenization?) will cause breakdown of the solid to form microcapsules (or micronized lamellar solid drug) in suspension.

General procedure:

I will first attempt to prepare a 1% prednisolone suspension using a 5% (w/w) solid dispersion of drug + lecithin (or egg phosphatides). This solid will be ^{2, 3, 4, 5} dispersed in an aqueous medium (water + mannitol). Call this formulation FORMULATION 1.

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J. S. Kijjo

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BOOK NO. 20813 - 60

TITLE pH-precipitation of Lactz Acid Prior to Homogenization
 From Page No. — PCA or Project # — Protocol # — Date —

Purpose: Note an idea. Use pH-induced precipitation of Intracmarole prior to homogenization to reduce particle size. Use lactic acid to solubilize the drug. (See Chem. Pharm. Bull., 46(8), 1333-1334 (1998)).

Lactz Acid → pH → Homogenization → Add → Stabilize → Store
 + Drug or other means change or other means or other mechanical means Surfactant(s) micro after no

To Page No. —

SIGNATURE

Mah J

DATE

REVIEW

[Signature]

DATE

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